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Multiple vectors effectively achieve gene transfer in a murine cardiac transplantation model. Immunosuppression with TGF-beta 1 or vIL-10.

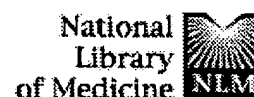
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Qin L, Chavin KD, Ding Y, Favaro JP, Woodward JE, Lin J, Tahara H, Robbins P, Shaked A, Ho DY, et al.

Department of Microbiology, Medical University of South Carolina, Charleston 29425, USA.

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The application of gene transfer techniques to organ transplantation offers the potential for modulation of immunity directly within an allograft without systemic side effects. Expression vectors and promoter elements are important determinants of gene transfer and expression. In this study, various vectors (naked plasmid DNA, retroviral vector, herpes simplex viral vector, and adenoviral vector) with various promoters (RSV-LTR, SV40, MuLV-LTR, HCMVie1) were directly compared to demonstrate the successful gene transfer and expression of beta-galactosidase in murine myoblasts in vitro and within murine heterotopic, nonvascularized cardiac isografts or allografts in vivo. Expression of transferred genes was not toxic to cells and strength of expression varied according to the type of vector. Plasmid DNA was expressed in myocytes, retroviral vector was expressed in the graft infiltrating cells, and herpes simplex and adenoviral vectors were expressed in both myocytes and graft-infiltrating cells. Preliminary studies evaluated the ability of these vectors to deliver immunologically important signals. Allografts injected with pSVTGF-beta 1, a plasmid-encoding transforming growth factor beta 1 (TGF-beta 1) under the control of the SV40 promoter, showed significant prolongation of graft survival of 26.3 +/- 2.5 days compared with 12.6 +/- 1.1 days for untreated allografts, and 12.5 +/- 1.5 days for the allografts injected with control plasmid ($P < 0.05$). Allografts injected with MFG-vIL-10, a retroviral vector encoding viral interleukin-10 under the control of the MuLV-LTR, showed prolongation of graft survival of 36.7 +/- 1.3 days versus 12.6 +/- 1.1 days for the untreated allograft, and 13.5 +/- 2.0 days for the allografts injected with control retroviral vector ($P < 0.001$). Both vectors were transcriptionally active in vivo and did not appear to have toxic effects. Gene therapy for



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Vasc Endovascular Surg. 2003 May-Jun;37(3):229-30. No abstract available.
PMID: 12799735 [PubMed - in process]

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J Vasc Surg. 2003 Jun;37(6):1294-300.
PMID: 12764278 [PubMed - indexed for MEDLINE]

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J Vasc Surg. 2003 Feb;37(2):301-6.
PMID: 12563199 [PubMed - indexed for MEDLINE]

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J Surg Res. 2002 Dec;108(2):198-202.
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J Surg Res. 2002 Dec;108(2):191-7.
PMID: 12505041 [PubMed - indexed for MEDLINE]


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PMID: 12478202 [PubMed - indexed for MEDLINE]


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Ann Vasc Surg. 2002 Nov;16(6):779-83. Epub 2002 Oct 31.
PMID: 12404042 [PubMed - indexed for MEDLINE]


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- ☐ 11: [Skelly CL, Meyerson SL, Curi MA, Desai TR, Bassiouny HS, McKinsey JF, Gewertz BL, Schwartz LB.](#) Related Articles, Links

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Ann Vasc Surg. 2002 Jan;16(1):95-101. Epub 2002 Jan 16.
PMID: 11904812 [PubMed - indexed for MEDLINE]

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 Prevention of restenosis by a herpes simplex virus mutant capable of controlled long-term expression in vascular tissue in vivo.
Gene Ther. 2001 Dec;8(24):1840-6.
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The effects of extremely low shear stress on cellular proliferation and neointimal thickening in the failing bypass graft.

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PMID: 11436071 [PubMed - indexed for MEDLINE]

- ☐ **16:** [Desai TR, Meyerson SL, Skelly CL, MacKenzie KS, Bassiouny HS, Katz D, McKinsey JF, Gewertz BL, Schwartz LB.](#) [Related Articles, Links](#)



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- ☐ **17:** [Moawad J, Meyerson SL, Refai D, Skelly CL, Leiden JM, Schwartz LB.](#) [Related Articles, Links](#)



Adenoviral-mediated gene transfer in human and animal vein grafts using clinically relevant exposure times, pressures, and viral concentrations.

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The hemodynamics of vein grafts: measurement and meaning.

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Effective hemodynamic diameter: an intrinsic property of vein grafts with predictive value for patency.

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- ☐ **20:** [Baldwin ZK, Meyerson SL, Skelly CL, McKinsey JF, Bassiouny HS, MacDonald RL, Gewertz BL, Schwartz LB.](#) [Related Articles, Links](#)



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Ann Vasc Surg. 2000 May;14(3):210-5.
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